

**Claim 6** was rejected under 35 USC § 112 as containing subject matter not described in the specification. The applicant disagrees, nevertheless has amended claim 6 to limit the scope of the claim to advance prosecution of the present case.

**Claims 17-25** were rejected under 35 USC § 112 as failing to provide enablement for treatment of diseases associated with an inappropriate immune response *in vivo*. The applicant disagrees, nevertheless has amended claims 17, and 19-25, and has canceled claim 18 to advance prosecution of the present case. Amended claims 17, and 19-25 are now directed towards treatment of an immunocompetent cell.

### **35 USC § 102**

**Claims 1-3, 7, 9, and 10** were rejected under 35 USC § 102(b) as being anticipated by Patel et al. (J. Mol. Biol. (1997) 272, 645-664). The applicant disagrees. Among other elements, amended claim 1, and claims 2,3, 7, 9, and 10 by virtue of their dependence on amended claim 1 require that the aptamer has a "...length of between about **12 and 22 nucleic acid units**, inclusive...", and that "...the aptamer **reduces CD28 expression in an activated human T-cell...**" None of these elements is taught by Patel et al. Consequently, claims 1-3, 7, 9, and 10 are not anticipated by Patel et al.

**Claims 1-6, 8, and 10** were rejected under 35 USC § 102(b) as being anticipated by Sharma et al. (Anticancer Res. (1996) 16, 61-70). The applicant disagrees. Again, amended claim 1, and claims 2-6, 8, and 10 by virtue of their dependence on amended claim 1 require that "...the aptamer **reduces CD28 expression in an activated human T-cell...**" This element is not taught by Sharma et al. Consequently, claim 1, and claims 2-6, 8, and 10 are not anticipated by Sharma et al.

**Claims 1-6, 8, and 10** were rejected under 35 USC § 102(b) as being anticipated by Smith and Feigon (Nature (1992) 356, 164-167). The applicant disagrees. Once more, amended claim 1, and claims 2-6, 8, and 10 by virtue of their dependence on amended claim 1 require that "...the aptamer **reduces CD28 expression in an activated human T-cell...**" This element is not taught by Smith and Feigon. Consequently, claim 1, and claims 2-6, 8, and 10 are not anticipated by Smith and Feigon.

**35 USC § 103**

Claims 1-5, 7-10, 11, 13, and 16 were rejected under the judicially created doctrine of obviousness-type double patenting. The applicant disagrees, nevertheless a terminal disclaimer over U.S. Pat. No. 5,932,556 to overcome the Examiner's rejection is being filed concurrently with the present response.

**Allowable Subject Matter**

The applicant acknowledges the Examiner's statement of allowability of claims 12, 14, and 15.

**ATTACHED MARKED-UP VERSION OF CHANGES**

Attached hereto is a marked-up version of the changes made to the claims by the current amendment. The attached page is captioned "VERSION WITH MARKINGS TO SHOW CHANGES MADE".

**REQUEST FOR ALLOWANCE**

Claims 1-4, 6-17, and 19-25 are pending in this application. The applicant requests allowance of all pending claims.

Respectfully submitted,

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By: \_\_\_\_\_



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## VERSIONS WITH MARKING TO SHOW CHANGES MADE

### In the Claims

1. (Amended) An aptamer having a length of between about 12 and 22 nucleic acid units, inclusive, and having a sequence which includes at least two G-rich regions selected from the group consisting of GGnG, GGGG, GnGG, nGGG and GGGn, where G is guanidine and n is any nucleotide, and wherein the aptamer reduces CD28 expression in an activated human T-cell.
2. (Amended) The aptamer of claim 1 wherein at least two of the at least two regions are separated by [less than] two to seven nucleotides, inclusive.
5. Canceled.
6. (Amended) The aptamer of claim [2] 1 wherein the aptamer competes for a nucleic acid binding site of [immune regulatory protein is selected from the group of] SP1 [, NFkB, EGR1 and AP2].
17. (Amended) A method of treating [modulating] an immunocompetent cell [system response in a patient], comprising administering to the [patient] cell an aptamer according to claim 1 at a concentration effective to reduce CD28 expression.
18. Canceled.
19. (Amended) The method of claim [18] 17 wherein the immune competent cell is in a patient suffering from [condition comprises] a graft vs host response.
20. (Amended) The method of claim [18] 17 wherein the immune competent cell is in a patient suffering from [condition comprises] an autoimmune disease.
21. (Amended) The method of claim 20 wherein the [condition] autoimmune disease comprises rheumatoid arthritis.
22. (Amended) The method of claim 20 wherein the [condition] autoimmune disease comprises multiple sclerosis.

23. (Amended) The method of claim 20 wherein the [condition] autoimmune disease comprises lupus erythematosus.

24. (Amended) The method of claim 20 wherein the [condition] autoimmune disease comprises insulin dependent diabetes mellitus.

25. (Amended) The method of claim 20 wherein the [condition] autoimmune disease comprises psoriasis.